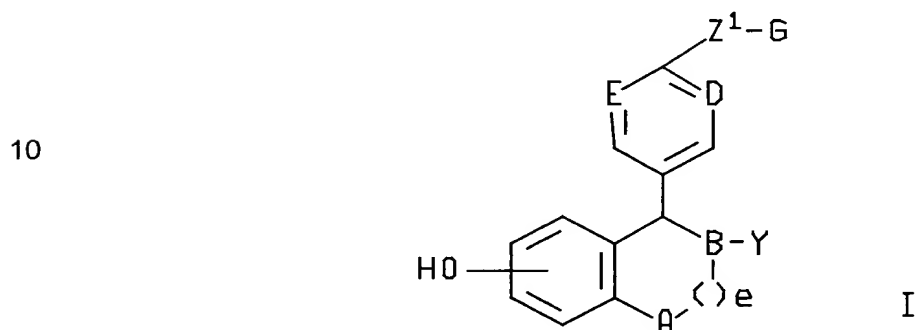


CLAIMS

What is claimed is:

1. A method of preventing breast cancer in a mammal which comprises
5 administering to a mammal in need of such prevention an effective amount of a compound of the formula I:



15 wherein:

A is selected from CH₂ and NR;

B, D and E are independently selected from CH and N;

Y is

- 20 (a) phenyl, optionally substituted with 1-3 substituents independently selected from R⁴;
- (b) naphthyl, optionally substituted with 1-3 substituents independently selected from R⁴;
- (c) C₃-C₈ cycloalkyl, optionally substituted with 1-2 substituents independently selected from R⁴;
- 25 (d) C₃-C₈ cycloalkenyl, optionally substituted with 1-2 substituents independently selected from R⁴;
- (e) a five membered heterocycle containing up to two heteroatoms selected from the group consisting of -O-, -NR²- and -S(O)_n-, optionally substituted with 1-3 substituents independently selected from R⁴;
- 30 (f) a six membered heterocycle containing up to two heteroatoms selected from the group consisting of -O-, -NR²- and -S(O)_n-, optionally substituted with 1-3 substituents independently selected from R⁴; or

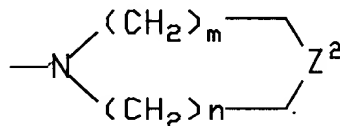
- (g) a bicyclic ring system consisting of a five or six membered heterocyclic ring fused to a phenyl ring, said heterocyclic ring containing up to two heteroatoms selected from the group consisting of -O-, -NR²-, NR²- and -S(O)_n-, optionally substituted with 1-3 substituents independently selected from R⁴;

Z¹ is

- (a) -(CH₂)_p W(CH₂)_q-;
 (b) -O(CH₂)_p CR⁵R⁶-;
 (c) -O(CH₂)_pW(CH₂)_q-;
 (d) -OCHR²CHR³-; or
 (e) -SCHR²CHR³-;

G is

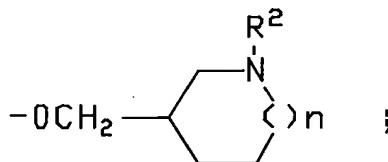
- (a) -NR⁷R⁸;
 (b)



wherein n is 0, 1 or 2; m is 1, 2 or 3; Z² is -NH-, -O-, -S-, or -CH₂-; optionally fused on adjacent carbon atoms with one or two phenyl rings and, optionally independently substituted on carbon with one to three substituents and, optionally, independently on nitrogen with a chemically suitable substituent selected from R⁴; or

- (c) a bicyclic amine containing five to twelve carbon atoms, either bridged or fused and optionally substituted with 1-3 substituents independently selected from R⁴;

Z¹ and G in combination may be



W is

5

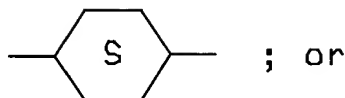
- (a) $-\text{CH}_2-$;
- (b) $-\text{CH}=\text{CH}-$;
- (c) $-\text{O}-$;
- (d) $-\text{NR}^2-$;
- (e) $-\text{S}(\text{O})_n-$;
- (f)

10



- (g) $-\text{CR}^2(\text{OH})-$;
- (h) $-\text{CONR}^2-$;
- (i) $-\text{NR}^2\text{CO}-$;
- (j)

15



- (k) $-\text{C}\equiv\text{C}-$;

R is hydrogen or $\text{C}_1\text{-C}_6$ alkyl;

20

R^2 and R^3 are independently

- (a) hydrogen; or
- (b) $\text{C}_1\text{-C}_4$ alkyl;

R^4 is

25

- (a) hydrogen;
- (b) halogen;
- (c) $\text{C}_1\text{-C}_6$ alkyl;
- (d) $\text{C}_1\text{-C}_4$ alkoxy;
- (e) $\text{C}_1\text{-C}_4$ acyloxy;
- (f) $\text{C}_1\text{-C}_4$ alkylthio;
- (g) $\text{C}_1\text{-C}_4$ alkylsulfinyl;
- (h) $\text{C}_1\text{-C}_4$ alkylsulfonyl;
- (i) hydroxy ($\text{C}_1\text{-C}_4$)alkyl;
- (j) aryl ($\text{C}_1\text{-C}_4$)alkyl;
- (k) $-\text{CO}_2\text{H}$;

30

- 5
- (l) -CN;
 - (m) -CONHOR;
 - (n) -SO₂NHR;
 - (o) -NH₂;
 - (p) C₁-C₄ alkylamino;
 - (q) C₁-C₄ dialkylamino;
 - (r) -NHSO₂R;
 - (s) -NO₂;
 - (t) -aryl; or
 - 10 (u) -OH.

R⁵ and R⁶ are independently C₁-C₈ alkyl or together form a C₃-C₁₀ carbocyclic ring;

R⁷ and R⁸ are independently

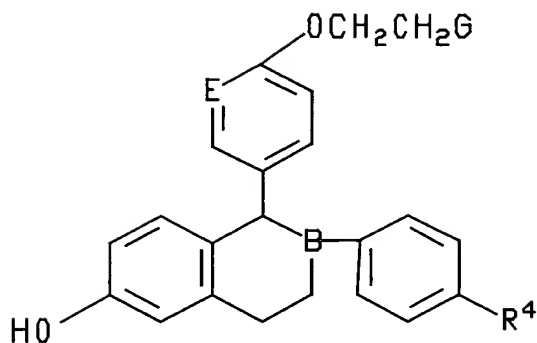
- 15
- (a) phenyl;
 - (b) a C₃-C₁₀ carbocyclic ring, saturated or unsaturated;
 - (c) a C₃-C₁₀ heterocyclic ring containing up to two heteroatoms, selected from -O-, -N- and -S-;
 - (d) H;
 - (e) C₁-C₆ alkyl; or
 - 20 (f) form a 3 to 8 membered nitrogen containing ring with R⁵ or R⁶;

R⁷ and R⁸ in either linear or ring form may optionally be substituted with up to three substituents independently selected from C₁-C₆ alkyl, halogen, alkoxy, hydroxy and carboxy;

- 25
- a ring formed by R⁷ and R⁸ may be optionally fused to a phenyl ring;
 - e is 0, 1 or 2;
 - m is 1, 2 or 3;
 - n is 0, 1 or 2;
 - p is 0, 1, 2 or 3;
 - q is 0, 1, 2 or 3;

30 and optical and geometric isomers thereof; and nontoxic pharmacologically acceptable acid addition salts, N-oxides, esters, and quaternary ammonium salts thereof.

2. A method of claim 1 wherein the compound is a compound of the formula:



5

3. A method of preventing breast cancer in a mammal which comprises administering to a mammal in need of such prevention an effective amount of (-)-*Cis*-6-phenyl-5-[4-(2-pyrrolidin-1-yl-ethoxy)-phenyl]-5,6,7,8-tetrahydro-naphthalen-2-ol, or a nontoxic pharmacologically acceptable acid addition salt, N-oxide, ester, or quaternary ammonium salt thereof.
- 10 4. A method of preventing breast cancer in a mammal which comprises administering to a mammal in need of such prevention an effective amount of *Cis*-6-phenyl-5-[4-(2-pyrrolidin-1-yl-ethoxy)-phenyl]-5,6,7,8-tetrahydro-naphthalen-2-ol, or an optical or geometric isomer thereof; or a nontoxic pharmacologically acceptable acid addition salt, N-oxide, ester, or quaternary ammonium salt thereof.
- 15